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(54) Title: COMPOSITIONS AND METHODS FOR DETECTING TARGET NUCLEIC ACID SEQUENCES UTILIZING FLANKING SEQUENCE ENZYME MOLECULES

(57) Abstract

Briefly stated, the present invention provides novel compositions and methods for detecting target nucleic acid sequences utilizing adjacent sequence-enzyme molecules. Within one aspect of the present invention, oligonucleotide-enzyme fusion molecules are provided, comprising an enzyme capable of cleaving scissile linkages and an oligonucleotide having the structure (NA₁)_x-S₂-(NA₂)_y wherein NA₁ and NA₂ are nucleic acid sequences, S is a scissile nucleic acid linkage, x, y, and z are integers from 1 to 1,000 and n is an integer from 1 to 10.

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Interns 11 Application No PCT/CA 94/00448

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| According to | o International Patent Classification (IPC) or to both national classi | fication and IPC | | |
| | SEARCHED | | · | |
| Minimum d IPC 6 | ocumentation searched (classification system followed by classification classification system followed by cl | tion symbols) | | |
| Documentat | ion searched other than minimum documentation to the extent that | such documents are included in the fields s | earched | |
| Electronic d | ata base consulted during the international search (name of data ba | se and, where practical, search terms used) | | |
| C. DOCUM | IENTS CONSIDERED TO BE RELEVANT | | | |
| Category * | Citation of document, with indication, where appropriate, of the r | elevant passages | Relevant to claim No. | |
| Y | WO,A,89 10415 (MEIOGENICS ,INC.) November 1989 cited in the application see the whole document | 2 | 1-14 | |
| Y | EP,A,O 461 731 (LONDON BIOTECHNO 18 December 1991 see page 15, line 5 - line 19 | LOGY LTD.) | 1-14 | |
| Y | SCIENCE, vol.110, 1988, LANCASTER, PA US Zuckermann, Ronald N. et al 'A h sequence-selective ribonuclease see the whole document | | 1-14 | |
| Y | EP,A,O 361 768 (SISKA DIAGNOSTIC April 1990 see abstract | S,INC.) 4 | 12,13 | |
| X Furt | ther documents are listed in the continuation of box C. | Patent family members are listed | in annex. | |
| 'A' docum consider a consider the consider the consider the consider the consideration of the consideration that the consideration the consideration the consideration that the conside | the defining the general state of the art which is not lered to be of particular relevance document but published on or after the international date ent which may throw doubts on priority claim(s) or is cited to establish the publication date of another on or other special reason (as specified) tent referring to an oral disclosure, use, exhibition or means ent published prior to the international filing date but han the priority date claimed actual completion of the international search | To later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention. 'X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone. 'Y' document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. '&' document member of the same patent family Date of mailing of the international search report | | |
| Name and | mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+ 31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+ 31-70) 340-3016 | Authorized officer Gurdjian, D | | |

INTERNATIONAL SEARCH REPORT

Internr 1 Application No
PCT/CA 94/00448

| ategory * | tion) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages | Relevant to claim No. | |
|-----------|---|-----------------------|--|
| | EP,A,O 361 983 (GENE TRAK SYSTEMS) 4 April 1990 see abstract | 1,14 | |
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INTERNATIONAL SEARCH REPORT

. .ormation on patent family members

Interny al Application No PCT/CA 94/00448

| Patent document cited in search report | Publication date | Patent family member(s) | Publication date |
|--|------------------|---|--|
| WO-A-8910415 | 02-11-89 | US-A- 50117 AU-A- 35697 EP-A- 03656 JP-T- 25041 | 789 24-11-89 663 02-05-90 |
| EP-A-0461731 | 18-12-91 | AT-T- 1084 CA-A- 12586 DE-A- 35861 DE-D- 35878 EP-A,B 01566 GB-A,B 21565 GB-A,B 21928 JP-A- 610013 US-A- 50574 US-A- 47456 | 22-08-89 183 16-07-92 383 18-08-94 541 02-10-85 518 09-10-85 389 27-01-88 398 07-01-86 412 15-10-91 |
| EP-A-0361768 | 04-04-90 | CA-A- 13333 JP-A- 22578 | |
| EP-A-0361983 | 04-04-90 | JP-A- 22578 | 898 18-10-90 |

Form PCT/ISA/210 (patent family annex) (July 1992)

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